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09/105/17

(FILE 'HOME' ENTERED AT 18:56:31 ON 19 OCT 2001)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, SCISEARCH' ENTERED AT 18:57:06 ON  
19 OCT 2001

L1 2893 S MICROBIAL (W) PRODUCTION  
L2 190 S L1 AND (AMINO (W) ACIDS)  
L3 28 S L2 AND CORYNEBACTERIUM  
L4 13 L3 AND LYSINE  
L5 0 L3 AND ((EXPORT) (W) (GENE OR CARRIER))  
L6 0 S L3 AND EXPORT (W) GENE  
L7 166 S EXPORT (W) GENE  
L8 0 S L3 AND L7  
L9 0 S L3 (P) L7  
L10 0 S L2 AND L7  
L11 0 S L2 AND EXPORT (W) GENE  
L12 61 S L7 AND MICROB?  
L13 1 S L12 AND CORYNEBACTERIUM  
L14 26 DUP REM L3 (2 DUPLICATES REMOVED)  
L15 13 DUP REM L4 (0 DUPLICATES REMOVED)

=> log off y

STN  
Search Strategy

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09/105/17

EAST

Search Strategy

Type	L #	Hits	Search Text	DBs	Time Stamp	Comments	Error Definition	Errors
1 BRS	L11	0	microbial adj production adj of adj amino adj acid	USPA T; US-P GPUB ; EPO; JPO; DERW ENT	2001/10/1 9 18:38			0
2 BRS	L7	702	microbial adj production	USPA T; US-P GPUB ; EPO; JPO; DERW ENT	2001/10/1 9 18:38			0
3 BRS	L13	314	L7 and (amino adj acid)	USPA T; US-P GPUB ; EPO; JPO; DERW ENT	2001/10/1 9 18:39			0
4 BRS	L19	47	L13 and Corynebacterium	USPA T; US-P GPUB ; EPO; JPO; DERW ENT	2001/10/1 9 18:40			0

Type	L #	Hits	Search Text	DBs	Time Stamp	Comments	Error Definition	Errorors
5	BRS	L25	0	L19 and (manufacturing adj L-lysine)	USPA T; US-P GPUB ; EPO; JPO; DERW ENT	2001/10/1 9 18:42		0
6	BRS	L37	1	L19 and ((export) adj (gene or carrier))	USPA T; US-P GPUB ; EPO; JPO; DERW ENT	2001/10/1 9 18:47		0
7	BRS	L31	11	L19 and lysine	USPA T; US-P GPUB ; EPO; JPO; DERW ENT	2001/10/1 9 18:48		0

155

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09/105117

## Connecting via Winsock to STN

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NEWS 1 Web Page URLs for STN Seminar Schedule - N. America  
NEWS 2 Dec 17 The CA Lexicon available in the CAPLUS and CA files  
NEWS 3 Feb 06 Engineering Information Encompass files have new names  
NEWS 4 Feb 16 TOXLINE no longer being updated  
NEWS 5 Apr 23 Search Derwent WPINDEX by chemical structure  
NEWS 6 Apr 23 PRE-1967 REFERENCES NOW SEARCHABLE IN CAPLUS AND CA  
NEWS 7 May 07 DGENE Reload  
NEWS 8 Jun 20 Published patent applications (A1) are now in USPATFULL  
NEWS 9 JUL 13 New SDI alert frequency now available in Derwent's DWPI and DPCI  
NEWS 10 Aug 23 In-process records and more frequent updates now in MEDLINE  
NEWS 11 Aug 23 PAGE IMAGES FOR 1947-1966 RECORDS IN CAPLUS AND CA  
NEWS 12 Aug 23 Adis Newsletters (ADISNEWS) now available on STN  
NEWS 13 Sep 17 IMSworld Pharmaceutical Company Directory name change to PHARMASEARCH  
NEWS 14 Oct 09 Korean abstracts now included in Derwent World Patents Index  
NEWS 15 Oct 09 Number of Derwent World Patents Index updates increased  
NEWS 16 Oct 15 Calculated properties now in the REGISTRY/ZREGISTRY File

NEWS EXPRESS August 15 CURRENT WINDOWS VERSION IS V6.0c,  
CURRENT MACINTOSH VERSION IS V6.0 (ENG) AND V6.0J (JP),  
AND CURRENT DISCOVER FILE IS DATED 07 AUGUST 2001

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NEWS INTER General Internet Information  
NEWS LOGIN Welcome Banner and News Items  
NEWS PHONE Direct Dial and Telecommunication Network Access to STN  
NEWS WWW CAS World Wide Web Site (general information)

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FILE 'HOME' ENTERED AT 18:56:31 ON 19 OCT 2001

```
=> file biosis caplus embase medline scisearch  
COST IN U.S. DOLLARS  
SINCE FILE  
ENTRY  
TOTAL  
SESSION  
0.15  
0.15  
FULL ESTIMATED COST
```

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FILE 'MEDLINE' ENTERED AT 18:57:06 ON 19 OCT 2001

FILE 'SCISEARCH' ENTERED AT 18:57:06 ON 19 OCT 2001  
COPYRIGHT (C) 2001 Institute for Scientific Information (ISI) (R)

```
=> s microbial (w) production  
L1 2893 MICROBIAL (W) PRODUCTION
```

```
=> s L1 and (amino (w) acids)  
L2 190 L1 AND (AMINO (W) ACIDS)
```

```
=> s L2 and Corynebacterium  
L3 28 L2 AND CORYNEBACTERIUM
```

```
=> L3 and lysine  
L4 13 L3 AND LYSINE
```

```
=> L3 and ((export) (w) (gene or carrier))  
L5 0 L3 AND ((EXPORT) (W) (GENE OR CARRIER))
```

```
=> s L3 and export (w) gene  
L6 0 L3 AND EXPORT (W) GENE
```

```
=> s export (w) gene  
L7 166 EXPORT (W) GENE
```

```
=> s L3 and L7  
L8 0 L3 AND L7
```

```
=> s L3 (p) L7  
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH  
FIELD CODE - 'AND' OPERATOR ASSUMED 'L13 (P) L37'  
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH  
FIELD CODE - 'AND' OPERATOR ASSUMED 'L14 (P) L38'  
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH  
FIELD CODE - 'AND' OPERATOR ASSUMED 'L15 (P) L39'  
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH  
FIELD CODE - 'AND' OPERATOR ASSUMED 'L16 (P) L40'  
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH  
FIELD CODE - 'AND' OPERATOR ASSUMED 'L17 (P) L41'  
L9 0 L3 (P) L7
```

=>  
=> s L2 and L7  
L10 0 L2 AND L7

=> s L2 and export (w) gene  
L11 0 L2 AND EXPORT (W) GENE

=> s L7 and microb?  
L12 61 L7 AND MICROB?

=> s L12 and Corynebacterium  
L13 1 L12 AND CORYNEBACTERIUM

=> dup rem L3  
PROCESSING COMPLETED FOR L3  
L14 26 DUP REM L3 (2 DUPLICATES REMOVED)

=> dup rem L4  
PROCESSING COMPLETED FOR L4  
L15 13 DUP REM L4 (0 DUPLICATES REMOVED)

=> dis L13 ibib kwic

L13 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2001 ACS  
ACCESSION NUMBER: 1997:475788 CAPLUS  
DOCUMENT NUMBER: 127:172154  
TITLE: L-Lysine export from **Corynebacterium**  
glutamicum. Physiological and molecular-biological  
characterization of the carrier-mediated export of a  
primary metabolite  
AUTHOR(S): Vrljic, Maria-Marina  
CORPORATE SOURCE: Inst. Biotechnologie, Forschungszentrum Juelich  
G.m.b.H., Juelich, D-52425, Germany  
SOURCE: Ber. Forschungszent. Juelich (1997), Juel-3349, 1-115  
pp.  
CODEN: FJBEE5; ISSN: 0366-0885  
DOCUMENT TYPE: Report  
LANGUAGE: German  
TI L-Lysine export from **Corynebacterium** glutamicum. Physiological  
and molecular-biological characterization of the carrier-mediated export  
of a primary metabolite  
AB The gene for the Lys-excretion carrier was isolated from *C. glutamicum*  
and  
the Lys export was analyzed physiol. A system was established which  
induces the Lys excretion in dependence of Met. The mutant NA8, defect  
in  
Lys export, was isolated. The L-Lys export (LysE) gene encodes a  
polypeptide of 236 amino acids with the potential to span the membrane 6  
times and a mol. wt. of 2,5425 Da. With overexpressed LysE, L-Lys was  
exported at a rate of 3.76 nmol/min/mg dry wt. which lead to a 10-fold  
increased Lys excretion rate. The LysG (governing L-Lys **export**)  
gene is localized immediately adjacent to LysE, but is  
transcribed divergently. The deduced polypeptide (290 amino acids) has  
a  
helix-turn-helix motive at the aminotermminus. At the sequence level,  
LysG shows  $\text{ltoreq.} 35\%$  identity to prokaryotic, autoregulatory transcriptional

regulators. LysG acts in trans and leads to a decrease of the Lys excretion by *C. glutamicum*. For the Lys-export defect mutant C. glutamicum NA8, the transition G1594.fwdarw.A1594 was shown which results in a stop-codon in the LysE gene. The resulting LysE polypeptide in C. glutamicum NA8 is shortened for 43 amino acids. The growth of a LysEG deletion mutant was abolished on a minimal medium in the presence of Lys-contg. dipeptides. The quantification of the intracellular L-Lys concns. revealed an accumulation of Lys .ltoreq.1,100 mM. The results suggest that the physiol. function of the Lys export carrier of C. glutamicum is to avoid extremely high intracellular Lys concns.

ST lysine excretion carrier ***Corynebacterium*** gene sequence; protein sequence ***Corynebacterium*** lysine excretion carrier

IT Amino acid transport (biological)  
(carrier-mediated, export; lysine export from ***Corynebacterium*** glutamicum, carrier-supported export of a primary metabolite)

IT Helix-turn-helix  
(gene lysG protein; lysine export from ***Corynebacterium*** glutamicum, carrier-supported export of a primary metabolite)

IT Proteins (specific proteins and subclasses)  
RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)  
(gene lysG, (governing lysine export); lysine export from ***Corynebacterium*** glutamicum, carrier-supported export of a primary metabolite)

IT Genes (**microbial**)  
RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)  
(lysE; lysine export from ***Corynebacterium*** glutamicum, carrier-supported export of a primary metabolite)

IT Genes (**microbial**)  
RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)  
(lysG (governing lysine export); lysine export from ***Corynebacterium*** glutamicum, carrier-supported export of a primary metabolite)

IT ***Corynebacterium*** glutamicum  
DNA sequences  
Protein sequences  
(lysine export from ***Corynebacterium*** glutamicum, carrier-supported export of a primary metabolite)

IT Amino acid transporters  
RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)  
(lysine-transporting, gene lySE; lysine export from ***Corynebacterium*** glutamicum, carrier-supported export of a primary metabolite)

IT 63-68-3, L-Methionine, biological studies  
RL: BAC (Biological activity or effector, except adverse); BIOL (Biological study)  
(induces lysine excretion; lysine export from ***Corynebacterium*** glutamicum, carrier-supported export of a primary metabolite)

IT 184922-77-8, GenBank X96471-derived protein GI 1729755  
RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)  
(lysine export from ***Corynebacterium*** glutamicum,

carrier-supported export of a primary metabolite)  
IT 184922-76-7, GenBank X96471-derived protein GI 1729754 184922-78-9,  
GenBank X96471-derived protein GI 1729756  
RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological  
study); OCCU (Occurrence)  
(lysine export from **Corynebacterium** glutamicum,  
carrier-supported export of a primary metabolite)  
IT 56-87-1, L-Lysine, biological studies  
RL: BPR (Biological process); BIOL (Biological study); PROC (Process)  
(lysine export from **Corynebacterium** glutamicum,  
carrier-supported export of a primary metabolite)  
IT 184343-19-9, GenBank X96471  
RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological  
study); OCCU (Occurrence)  
(nucleotide sequence; lysine export from **Corynebacterium**  
glutamicum, carrier-supported export of a primary metabolite)

=> dis L14 1-26 ibib kwic

L14 ANSWER 1 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS  
ACCESSION NUMBER: 2001:314459 BIOSIS  
DOCUMENT NUMBER: PREV200100314459  
TITLE: Effect of gluconic acid as a secondary carbon source on  
non-growing L-lysine producers cells of  
**Corynebacterium** glutamicum. Purification and  
properties of 6-phosphogluconate dehydrogenase.  
AUTHOR(S): Bianchi, Daniella; Bertrand, Olivier; Haupt, Karsten;  
Coello, Nereida (1)  
CORPORATE SOURCE: (1) Instituto de Biología Experimental, Universidad  
Central  
Venezuela deVenezuela, Caracas, 1041-A: ncoello@uole.com.ve  
SOURCE: Enzyme and Microbial Technology, (June 7, 2001) Vol. 28,  
No. 9-10, pp. 754-759. print.  
ISSN: 0141-0229.  
DOCUMENT TYPE: Article  
LANGUAGE: English  
SUMMARY LANGUAGE: English  
TI Effect of gluconic acid as a secondary carbon source on non-growing  
L-lysine producers cells of **Corynebacterium** glutamicum.  
Purification and properties of 6-phosphogluconate dehydrogenase.  
AB We studied the production of L-lysine in **Corynebacterium**  
glutamicum ATCC 21543 non growing cells obtained by nutrient limitation.  
Statistical analysis revealed significant differences in the L-lysine  
titers of. . .  
IT . . .  
Engineering; Methods and Techniques; Nutrition  
IT Chemicals & Biochemicals  
6-phosphogluconate dehydrogenase: amino acid sequence, analysis,  
molecular properties, pH, purification; L-lysine: microbial  
production, yield; amino acids: analysis;  
carbon sources; gluconic acid: secondary carbon source  
ORGN . . .  
Microorganisms; Irregular Nonsporing Gram-Positive Rods: Actinomycetes  
and Related Organisms, Eubacteria, Bacteria, Microorganisms;  
Microorganisms  
ORGN Organism Name

Bacillus subtilis (Endospore-forming Gram-Positives);  
**Corynebacterium** glutamicum (Irregular Nonsporing Gram-Positive Rods): non-growing cells; Escherichia coli (Enterobacteriaceae); bacteria (Bacteria); microorganisms (Microorganisms)

ORGN Organism Superterms

Bacteria; Eubacteria; Microorganisms

L14 ANSWER 2 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS  
ACCESSION NUMBER: 2001:421002 BIOSIS  
DOCUMENT NUMBER: PREV200100421002  
TITLE: L-glutamate fermentation and metabolic engineering:  
Studies  
on the L-glutamate production mechanism in Coryneform  
bacteria.  
AUTHOR(S): Nakamatsu, Tsuyoshi  
SOURCE: Nippon Nogeikagaku Kaishi, (Jun., 2001) Vol. 75, No. 6,  
pp.  
683-686. print.  
ISSN: 0002-1407.

DOCUMENT TYPE: General Review

LANGUAGE: Japanese

SUMMARY LANGUAGE: English

IT Major Concepts

Biochemistry and Molecular Biophysics; Bioprocess Engineering;  
Metabolism

IT Chemicals & Biochemicals

**amino acids**: large-scale **microbial**  
**production**; glutamate: large-scale **microbial**  
**production**; oxoglutarate dehydrogenase

ORGN Super Taxa

Irregular Nonsporing Gram-Positive Rods: Actinomycetes and Related  
Organisms, Eubacteria, Bacteria, Microorganisms

ORGN Organism Name

**Corynebacterium** spp. (Irregular Nonsporing Gram-Positive  
Rods)

ORGN Organism Superterms

Bacteria; Eubacteria; Microorganisms

L14 ANSWER 3 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS

ACCESSION NUMBER: 2001:174925 BIOSIS

DOCUMENT NUMBER: PREV200100174925

TITLE: MALDI-TOF MS for quantification of substrates and products  
in cultivations of **Corynebacterium** glutamicum.

AUTHOR(S): Wittmann, Christoph (1); Heinze, Elmar

CORPORATE SOURCE: (1) Biochemical Engineering Institute, Saarland  
University,

66041, Saarbruecken: c.wittmann@rz.uni-sb.de Germany

SOURCE: Biotechnology and Bioengineering, (March 20, 2001) Vol.  
72,

No. 6, pp. 642-647. print.

ISSN: 0006-3592.

DOCUMENT TYPE: Article

LANGUAGE: English

SUMMARY LANGUAGE: English

TI MALDI-TOF MS for quantification of substrates and products in  
cultivations

of **Corynebacterium** glutamicum.

IT Major Concepts

Biochemistry and Molecular Biophysics; Bioprocess Engineering; Methods and Techniques

IT Chemicals & Biochemicals

**amino acids: microbial production**

, quantitative analysis; products: quantitative analysis; substrates: quantitative analysis

L14 ANSWER 4 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:761605 CAPLUS

DOCUMENT NUMBER: 134:99608

TITLE: Development and use of miniaturized parallel experiment technology for bioprocess development

AUTHOR(S): Altenbach-Rehm, Jutta

CORPORATE SOURCE: Institut fur Biotechnologie, Julich, JUL-3782, Germany

SOURCE: Ber. Forschungszent. Juelich (2000), Juel-3782, a-f, i-iv, 1-233

CODEN: FJBEE5; ISSN: 0366-0885

DOCUMENT TYPE: Report

LANGUAGE: German

AB The fed-batch technique is nowadays the std. operation mode for high performance **microbial prodn.** processes. Shake flasks are widely used a simple bioreactors in batch process development. Because of uncontrolled changes in pH and reduced oxygen transfer rates parallel operated shake flasks can usually not be applied for microbial fed-batch process development. Due to the need to operate controlled stirred tank reactors the development of specific fermn. strategies is expensive and time consuming. To address these issues a miniature fed-batch technique was developed in cooperation with DASGIP mbH, Julich, and INFORS AG, Basel. Controlled batch and fed-batch fermns. can be performed in 16 parallel small scale bioreactors. The new technique allows the feeding of up to 4 different substrates and parallel pH control

based on user-defined profiles. The feeding assembly is sterilized chem. using dimethylcarbonate. To overcome the limitations of shake flasks, small scale bubble columns were developed. Gas distribution is performed with a new type of sterile sparger. Transferring fed-batch fermns. from small scale bubble columns to a stirred tank reactor a scale up factor of 20 was achieved. Escherichia coli K12 was chosen to test the new parallel bioreactor technique. Compared to shake flask fermns. the cell concn. was

50% higher due to efficient oxygen transfer. In fed-batch fermns. with pH-controlled substrate feeding up to 35 g/l DCW were achieved. For the 1st time, a parallel optimization of feeding profiles in parallel small scale fed-batch expts. with successful scale-up to a lab. bioreactor was performed. Escherichia coli BL 21 (DE3) pLySS produces the recombinant enzyme GDP-.alpha.-D-mannose-pyrophosphorylase after induction with isopropyl-.beta.-D-thiogalactoside (IPTG). Single induction with 0,5 mM IPTG resulted in a low specific enzyme activity of 1,6 U/g DCW (dry cell wt.). For the optimization of enzyme expression a genetic algorithm was used. A final enzyme activity of 111 U/g DCW was achieved for optimal substrate and inducer feeding profiles. To demonstrate the advantages of this new parallel bioreactor technique different strains of industrial relevance were investigated. **Corynebacterium glutamicum**, **Staphylococcus carnosus** and **Ashbya gossypii**.

ST bioreactor bubble column fed batch fermn; GDP mannose pyrophosphorylase bubble column fed batch; Staphylococcus bubble column fed batch fermn;

isoleucine bubble column fed batch **Corynebacterium**; riboflavin  
 bubble column fed batch Ashbya  
 IT **Amino acids**, biological studies  
 RL: BPR (Biological process); BIOL (Biological study); PROC (Process)  
     (amino acid consumption in riboflavin prodn. by Ashbya gossypii in  
     parallel bubble columns with fed-batch technique)  
 IT **Corynebacterium glutamicum**  
     (L-isoleucine prodn. by **Corynebacterium glutamicum** in  
     parallel bubble columns with fed-batch technique)  
 IT 73-32-5P, L-Isoleucine, biological studies  
 RL: BMF (Bioindustrial manufacture); BPR (Biological process); BIOL  
     (Biological study); PREP (Preparation); PROC (Process)  
     (L-isoleucine prodn. by **Corynebacterium glutamicum** in  
     parallel bubble columns with fed-batch technique, amino acid  
     consumption in riboflavin prodn. by Ashbya gossypii)  
 IT 61-90-5, L-Leucine, biological studies  
 RL: BPR (Biological process); MFM (Metabolic formation); BIOL (Biological  
     study); FORM (Formation, nonpreparative); PROC (Process)  
     (L-isoleucine prodn. by **Corynebacterium glutamicum** in  
     parallel bubble columns with fed-batch technique, amino acid  
     consumption in riboflavin prodn. by Ashbya gossypii)

L14 ANSWER 5 OF 26 CAPLUS COPYRIGHT 2001 ACS  
 ACCESSION NUMBER: 1999:244776 CAPLUS  
 DOCUMENT NUMBER: 130:266420  
 TITLE: Method for **microbial production of amino acids** of the aspartate and/or glutamate family and agents which can be used in said method  
 INVENTOR(S): Eikmanns, Bernd; Peters-Wendisch, Petra; Sahm, Hermann  
 PATENT ASSIGNEE(S): Forschungszentrum Julich G.m.b.H., Germany  
 SOURCE: PCT Int. Appl., 40 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: German  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9918228	A2	19990415	WO 1998-EP6210	19980930
WO 9918228	A3	19990520		
W: AU, BR, CA, CN, HU, ID, JP, KR, MX, RU, SK, US				
RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
DE 199831609	A1	19990415	DE 1998-199831609	19980714
AU 9911482	A1	19990427	AU 1999-11482	19980930
EP 1015621	A2	20000705	EP 1998-954301	19980930
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
BR 9813021	A	20000815	BR 1998-13021	19980930
PRIORITY APPLN. INFO.:			DE 1997-19743894 A	19971004
			DE 1998-199831609 A	19980714
			WO 1998-EP6210 W	19980930

TI Method for **microbial production of amino acids** of the aspartate and/or glutamate family and agents which can be used in said method

AB The invention relates to a method for **microbial prodn.** of **amino acids** of the aspartate and/or glutamate family in which the pyruvate carboxylase activity is increased by genetically changing the enzyme and/or the pyruvate carboxylase gene expression of a microorganism which produces the corresponding amino acid.  
In addn., the invention relates to a pyruvate carboxylase gene and addnl. agents which can be used in the inventive method.

ST amino acid fermn **Corynebacterium** pyruvate carboxylase genetic engineering

IT **Corynebacterium glutamicum**  
Fermentation  
(**microbial prodn. of amino acids**  
of the aspartate and/or glutamate family and agents which can be used in said method)

IT **Amino acids**, preparation  
RL: BMF (Bioindustrial manufature); BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(**microbial prodn. of amino acids**  
of the aspartate and/or glutamate family and agents which can be used in said method)

IT Genetic engineering  
(**microbial prodn. of amino acids**  
of the aspartate and/or glutamate family and modification of **Corynebacterium pyc** gene in said method)

IT Genes (microbial)  
RL: BPR (Biological process); BIOL (Biological study); PROC (Process)  
(**pyc; microbial prodn. of amino acids** of the aspartate and/or glutamate family and modification of **Corynebacterium pyc** gene in said method)

IT 56-86-0P, L-Glutamic acid, preparation 56-87-1P, L-Lysine, preparation 72-19-5P, L-Threonine, preparation 74-79-3P, L-Arginine, preparation 672-15-1P, L-Homoserine  
RL: BMF (Bioindustrial manufature); BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(**microbial prodn. of amino acids**  
of the aspartate and/or glutamate family and agents which can be used in said method)

IT 9014-19-1, Pyruvate carboxylase 204116-18-7 208541-81-5  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(**microbial prodn. of amino acids**  
of the aspartate and/or glutamate family and agents which can be used in said method)

L14 ANSWER 6 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS  
ACCESSION NUMBER: 1999:175414 BIOSIS  
DOCUMENT NUMBER: PREV199900175414  
TITLE: Cloning of the transketolase gene and the effect of its dosage on aromatic amino acid production in **Corynebacterium glutamicum**.  
AUTHOR(S): Ikeda, M. (1); Okamoto, K.; Katsumata, R.  
CORPORATE SOURCE: (1) Technical Research Laboratories, Kyowa Hakko Kogyo Co., Ltd., Hofu, Yamaguchi, 747-8522 Japan  
SOURCE: Applied Microbiology and Biotechnology, (Feb., 1999) Vol. 51, No. 2, pp. 201-206.  
ISSN: 0175-7598.  
DOCUMENT TYPE: Article

LANGUAGE: English  
 TI Cloning of the transketolase gene and the effect of its dosage on aromatic amino acid production in **Corynebacterium glutamicum**.  
 AB. . . enzyme of the non-oxidative pentose phosphate pathway. The effect of its overexpression on aromatic amino acid production was investigated in **Corynebacterium glutamicum**, a typical amino-acid-producing organism. For this purpose, the transketolase gene of the organism was cloned on the basis of. . . the presence of the gene in high copy numbers enabled tyrosine, phenylalanine and tryptophan producers to accumulate 5%-20% more aromatic amino acids. These results indicate that overexpressed transketolase activity operates to redirect the glycolytic intermediates toward the nonoxidative pentose phosphate pathway in. . .  
 IT Major Concepts  
     Bioprocess Engineering; Enzymology (Biochemistry and Molecular Biophysics); Molecular Genetics (Biochemistry and Molecular Biophysics)  
 IT Chemicals & Biochemicals  
     aromatic amino acids: microbial production; transketolase [EC 2.2.1.1]; **Corynebacterium** transketolase gene (Irregular Nonsporing Gram-Positive Rods)

L14 ANSWER 7 OF 26 CAPLUS COPYRIGHT 2001 ACS  
 ACCESSION NUMBER: 1998:277651 CAPLUS  
 DOCUMENT NUMBER: 128:307587  
 TITLE: **Microbial production of substances from aromatic metabolism**  
 INVENTOR(S): Sprenger, Georg; Siewe, Ruth; Sahm, Hermann; Karutz, Martin; Sonke, Theodorus  
 PATENT ASSIGNEE(S): Forschungszentrum Juelich G.m.b.H., Germany; Holland Sweetener Co. V.o.F.  
 SOURCE: Ger. Offen., 14 pp.  
 CODEN: GWXXBX  
 DOCUMENT TYPE: Patent  
 LANGUAGE: German  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
DE 19644566	A1	19980430	DE 1996-19644566	19961026
WO 9818936	A1	19980507	WO 1997-NL582	19971017
W:	AL, AU, BA, BB, BG, BR, CA, CN, CU, CZ, EE, GE, HU, ID, IL, IS, JP, KP, KR, LC, LK, LR, LT, LV, MG, MK, MN, MX, NO, NZ, PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG			
AU 9747277	A1	19980522	AU 1997-47277	19971017
EP 934418	A1	19990811	EP 1997-909748	19971017
R:	AT, BE, CH, DE, DK, ES, FR, GB, IT, LI, NL, SE, PT, FI			
CN 1241214	A	20000112	CN 1997-180908	19971017
JP 2001506486	T2	20010522	JP 1998-520318	19971017
PRIORITY APPLN. INFO.:			DE 1996-19644566 A	19961026
			WO 1997-NL582	W 19971017

TI Microbial production of substances from aromatic metabolism

IT Transport proteins  
RL: BOC (Biological occurrence); BPN (Biosynthetic preparation); BUU (Biological use, unclassified); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); USES (Uses)  
(gene glf glucose facilitator protein, of Zymomonas mobilis;  
**microbial prodn.** of substances from arom. metab.)

IT Genes (microbial)  
RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)  
(glf, for glucose facilitator protein of Zymomonas mobilis;  
**microbial prodn.** of substances from arom. metab.)

IT Genes (microbial)  
RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)  
(glk, for glucokinase of Zymomonas mobilis; **microbial prodn.** of substances from arom. metab.)

IT Pentose phosphate pathway  
(intermediates of, in amino acid manuf.; **microbial prodn.** of substances from arom. metab.)

IT Bacillus (bacterium genus)

Brevibacterium

**Corynebacterium**

Escherichia

Escherichia coli

Fermentation

Microorganism

Molecular cloning

Serratia

(**microbial prodn.** of substances from arom. metab.)

IT Transport proteins  
RL: BOC (Biological occurrence); BPN (Biosynthetic preparation); BUU (Biological use, unclassified); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); USES (Uses)  
(**microbial prodn.** of substances from arom. metab.)

IT Amino acids, preparation  
RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(**microbial prodn.** of substances from arom. metab.)

IT Plasmids  
(pZ4557tal; **microbial prodn.** of substances from arom. metab.)

IT Plasmids  
(pZ4557tkt; **microbial prodn.** of substances from arom. metab.)

IT Plasmids  
(pZ4557tkttal; **microbial prodn.** of substances from arom. metab.)

IT Genes (microbial)  
RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)  
(talB; **microbial prodn.** of substances from arom. metab.)

IT Genes (microbial)  
RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)  
(tktA; **microbial prodn.** of substances from arom.

metab.)  
 IT 9001-36-9P, Glucokinase  
 RL: BOC (Biological occurrence); BPN (Biosynthetic preparation); BUU (Biological use, unclassified); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); USES (Uses)  
     (gene glk, of Zymomonas mobilis; **microbial prodn.**  
     of substances from arom. metab.)  
 IT 585-18-2, Erythrose-4-phosphate  
 RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)  
     (in amino acid manuf.; **microbial prodn.** of  
     substances from arom. metab.)  
 IT 9014-46-4P, Transaldolase 9014-48-6P, Transketolase  
 RL: BOC (Biological occurrence); BPN (Biosynthetic preparation); BUU (Biological use, unclassified); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); USES (Uses)  
     (**microbial prodn.** of substances from arom. metab.)  
 IT 63-91-2P, L-Phenylalanine, preparation  
 RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
     (**microbial prodn.** of substances from arom. metab.)

L14 ANSWER 8 OF 26 CAPLUS COPYRIGHT 2001 ACS  
 ACCESSION NUMBER: 1995:674124 CAPLUS  
 DOCUMENT NUMBER: 123:54314  
 TITLE: Enhancement of reduced NADP production for enhanced  
       **microbial production** of biochemicals  
 INVENTOR(S): Kojima, Hiroyuki; Totsuka, Kazuhiko  
 PATENT ASSIGNEE(S): Ajinomoto Co., Inc., Japan  
 SOURCE: PCT Int. Appl., 32 pp.  
       CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: Japanese  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9511985	A1	19950504	WO 1994-JP1791	19941026
W: AU, BR, CA, CN, CZ, HU, JP, KR, PL, RU, SK, US, VN				
RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
CA 2175042	AA	19950504	CA 1994-2175042	19941026
AU 9480026	A1	19950522	AU 1994-80026	19941026
AU 687458	B2	19980226		
EP 733712	A1	19960925	EP 1994-931158	19941026
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT,				
SE				
BR 9407907	A	19961126	BR 1994-7907	19941026
HU 74840	A2	19970228	HU 1996-1085	19941026
ZA 9503350	A	19961025	ZA 1995-3350	19950425
US 5830716	A	19981103	US 1996-619521	19960429
CN 1139956	A	19970108	CN 1994-194707	19961026
PRIORITY APPLN. INFO.:			JP 1993-270828	19931028
			WO 1994-JP1791	19941026
TI	Enhancement of reduced NADP production for enhanced <b>microbial</b> <b>production</b> of biochemicals			
AB	The productivity of such substances as <b>L-amino acids</b> , antibiotics, vitamins, growth factors and physiol. active substances in			

the fermn. using a microorganism is improved by improving the productivity  
of reduced NADP in the cells of the microorganisms. Construction of pMW::THY contg. the Escherichia coli transhydrogenase gene, and introduction of the plasmid into the L-threonine-producing Escherichia coli B-3996 were shown. The recombinant E. coli B-3996 produced L-threonine .apprx.10% higher than did the parental strain.

IT **Corynebacterium glutamicum**  
Escherichia coli  
Fermentation  
(enhancement of reduced NADP prodn. for enhanced **microbial prodn. of biochms.**)

IT **Amino acids, preparation**  
RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(enhancement of reduced NADP prodn. for enhanced **microbial prodn. of biochms.**)

IT Plasmid and Episome  
(pHSG::THY; enhancement of reduced NADP prodn. for enhanced **microbial prodn. of biochms.**)

IT Plasmid and Episome  
(pMW::THY; enhancement of reduced NADP prodn. for enhanced **microbial prodn. of biochms.**)

IT Plasmid and Episome  
(pSU::THY; enhancement of reduced NADP prodn. for enhanced **microbial prodn. of biochms.**)

IT 9014-18-0, Nicotinamide nucleotide transhydrogenase  
RL: BAC (Biological activity or effector, except adverse); BIOL (Biological study)  
(enhancement of reduced NADP prodn. for enhanced **microbial prodn. of biochms.**)

IT 56-86-0P, L-Glutamic acid, preparation 56-87-1P, L-Lysine, preparation 61-90-5P, L-Leucine, preparation 63-91-2P, L-Phenylalanine, preparation 72-18-4P, L-Valine, preparation 72-19-5P, L-Threonine, preparation 73-32-5P, L-Isoleucine, preparation  
RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(enhancement of reduced NADP prodn. for enhanced **microbial prodn. of biochms.**)

IT 53-59-8P, NADP  
RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(reduced; enhancement of reduced NADP prodn. for enhanced **microbial prodn. of biochms.**)

L14 ANSWER 9 OF 26 SCISEARCH COPYRIGHT 2001 ISI (R)  
ACCESSION NUMBER: 95:184306 SCISEARCH  
THE GENUINE ARTICLE: QK574  
TITLE: METABOLIC DESIGN IN AMINO-ACID PRODUCING BACTERIUM  
**CORYNEBACTERIUM-GLUTAMICUM**  
AUTHOR: SAHM H (Reprint); EGELING L; EIKMANNS B; KRAMER R  
CORPORATE SOURCE: KFA JULICH GMBH, FORSCHUNGSZENTRUM, INST BIOTECHNOL,  
D-52425 JULICH, GERMANY (Reprint)  
COUNTRY OF AUTHOR: GERMANY  
SOURCE: FEMS MICROBIOLOGY REVIEWS, (FEB 1995) Vol. 16, No. 2-3,  
pp. 243-252.  
ISSN: 0168-6445.  
DOCUMENT TYPE: Article; Journal

FILE SEGMENT: LIFE  
LANGUAGE: ENGLISH  
REFERENCE COUNT: 36

\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

TI METABOLIC DESIGN IN AMINO-ACID PRODUCING BACTERIUM **CORYNEBACTERIUM-GLUTAMICUM**

AB The Gram-positive bacterium **Corynebacterium glutamicum** is used for the industrial production of **amino acids**, e.g. of L-glutamate and L-lysine. In the last 10 years, genetic engineering and amplification of relevant structural genes have become.

ST Author Keywords: **CORYNEBACTERIUM GLUTAMICUM; AMINO ACID PRODUCTION; METABOLIC DESIGN; L-LYSINE; L-THREONINE; L-ISOLEUCINE**  
STP KeyWords Plus (R): **L-THREONINE; L-LYSINE; BREVIBACTERIUM-LACTOFERMENTUM; HOMOSERINE DEHYDROGENASE; MICROBIAL PRODUCTION; RESISTANT MUTANTS; SPLIT PATHWAY; BIOSYNTHESIS; ISOLEUCINE; GENES**

L14 ANSWER 10 OF 26 SCISEARCH COPYRIGHT 2001 ISI (R)

ACCESSION NUMBER: 96:7187 SCISEARCH

THE GENUINE ARTICLE: TJ545

TITLE: METABOLIC DESIGN IN THE AMINO-ACID-PRODUCING BACTERIUM **CORYNEBACTERIUM-GLUTAMICUM**

AUTHOR: SAHM H (Reprint)

CORPORATE SOURCE: KFA JULICH GMBH, FORSCHUNGSZENTRUM, INST BIOTECHNOL, D-52425 JULICH, GERMANY (Reprint)

COUNTRY OF AUTHOR: GERMANY

SOURCE: FOLIA MICROBIOLOGICA, (1995) Vol. 40, No. 1, pp. 23-30.  
ISSN: 0015-5632.

DOCUMENT TYPE: Article; Journal

FILE SEGMENT: LIFE; AGRI

LANGUAGE: ENGLISH

REFERENCE COUNT: 26

\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

TI METABOLIC DESIGN IN THE AMINO-ACID-PRODUCING BACTERIUM **CORYNEBACTERIUM-GLUTAMICUM**

AB The Gram-positive bacterium **Corynebacterium glutamicum** is used for the industrial production of **amino acids**, e.g. of L-glutamate and L-lysine. By cloning and expressing the various genes of the L-lysine pathway in C. glutamicum we. . .

STP KeyWords Plus (R): **L-THREONINE; BREVIBACTERIUM-LACTOFERMENTUM; HOMOSERINE DEHYDROGENASE; MICROBIAL PRODUCTION; LYSINE BIOSYNTHESIS; RESISTANT MUTANTS; SPLIT PATHWAY; GENES; AMPLIFICATION; FERMENTATION**

L14 ANSWER 11 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1985:436156 CAPLUS

DOCUMENT NUMBER: 103:36156

TITLE: Optimization of amino acid production by automatic self-tuning digital control of redox potential  
AUTHOR(S): Radjai, Mohammad K.; Hatch, Randolph T.; Cadman, Theodore W.

CORPORATE SOURCE: Dep. Chem. Nucl. Eng., Univ. Maryland, College Park, MD, 20742, USA

SOURCE: Biotechnol. Bioeng. Symp. (1984), 14 (Symp. Biotechnol.)

Fuels Chem., 6th), 657-79

CODEN: BIBSBR; ISSN: 0572-6565

DOCUMENT TYPE: Journal  
LANGUAGE: English  
AB The microbial prodn. of homoserine [672-15-1], lysine [56-87-1], and valine [72-18-4] by an auxotrophic mutant of **Corynebacterium glutamicum** was investigated in a 16-L batch fermentor. Closed-loop digital control of redox potential was implemented using proportional-integral (PI) control of agitation rates. Due to the nonlinearity of the system, the PI controller parameters had to be changed during the course of the fermns. An automatic, self-tuning algorithm was developed for stable control of redox potential. This permitted exptl. optimization of total and selective amino acid prodn. Total amino acid yields of 35% from glucose were achieved compared to 23% reported in the literature for the same fermn.  
IT **Corynebacterium glutamicum**  
(amino acid manuf. with, optimization and redox potential control in)  
IT **Amino acids, preparation**  
RL: BMF (Bioindustrial manufacture); BIOL (Biological study); PREP (Preparation)  
(manuf. of, by fermn.)

L14 ANSWER 12 OF 26 CAPLUS COPYRIGHT 2001 ACS DUPLICATE 1  
ACCESSION NUMBER: 1979:20814 CAPLUS  
DOCUMENT NUMBER: 90:20814  
TITLE: **Microbial production of essential amino acids with Corynebacterium glutamicum mutants**  
AUTHOR(S): Nakayama, Kiyoshi; Araki, Kazumi; Kase, Hiroshi  
CORPORATE SOURCE: Tokyo Res. Lab., Kyowa Hakko Kogyo Co., Ltd.,  
Machida,  
Japan  
SOURCE: Adv. Exp. Med. Biol. (1978), 105(Nutr. Improv. Food Feed Proteins), 649-61  
CODEN: AEMBAP; ISSN: 0065-2598  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
TI **Microbial production of essential amino acids with Corynebacterium glutamicum mutants**  
AB **Amino acids** produced by microbial processes are generally L-forms. The stereospecificity of the **amino acids** produced by fermn. makes the process advantageous compared with synthetic processes. Microorganisms employed in microbial processes for amino acid prodn. are divided into 4 classes: wild-type, auxotrophic mutant, regulatory mutant, and auxotrophic regulatory mutant. Using such mutants of **Corynebacterium glutamicum**, all the essential **amino acids** but L-methionine are now being produced by direct fermn. from cheap C sources such as carbohydrate materials or acetic acid.  
ST amino acid manuf **Corynebacterium**  
IT **Corynebacterium glutamicum**  
(amino acid manuf. by)  
IT Fermentation  
(amino acids, by **Corynebacterium glutamicum**)  
IT **Amino acids, preparation**  
RL: BMF (Bioindustrial manufacture); BIOL (Biological study); PREP (Preparation)

(manuf. of, from carbohydrates by **Corynebacterium glutamicum**)

L14 ANSWER 13 OF 26 CAPLUS COPYRIGHT 2001 ACS  
ACCESSION NUMBER: 1976:521806 CAPLUS  
DOCUMENT NUMBER: 85:121806  
TITLE: **Microbial production of amino acid**  
INVENTOR(S): Tsuchida, Takayasu; Yoshihara, Yasuhiko; Kubota,  
Koji;  
Hirose, Yoshio  
PATENT ASSIGNEE(S): Ajinomoto Co., Inc., Japan  
SOURCE: Japan. Kokai, 5 pp.  
CODEN: JKXXAF  
DOCUMENT TYPE: Patent  
LANGUAGE: Japanese  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 51061690	A2	19760528	JP 1974-134879	19741122
TI	<b>Microbial production of amino acid</b>			
ST	amino acid manuf Brevibacterium; <b>Corynebacterium</b> amino acid manuf			
IT	Brevibacterium <b>Corynebacterium</b> (amino acid manuf. by)			
IT	Fermentation (amino acids, by <b>Corynebacterium</b> or Brevibacterium)			
IT	56-45-1P, preparation	73-22-3P, preparation		
	RL: PREP (Preparation)			
	(by fermn., with <b>Corynebacterium</b> )			

L14 ANSWER 14 OF 26 CAPLUS COPYRIGHT 2001 ACS  
ACCESSION NUMBER: 1975:529947 CAPLUS  
DOCUMENT NUMBER: 83:129947  
TITLE: **Microbial production of  
amino acids. VI. Formation of L-**  
**amino acids from**  
**DL-.alpha.-hydroxycarboxylic acids**  
AUTHOR(S): Matsushima, Hirochika; Murata, Keijiro; Mase, Yasuo  
CORPORATE SOURCE: Ferment. Res. Lab., Sankyo Co., Ltd., Tanashi, Japan  
SOURCE: Hakko Kogaku Zasshi (1975), 53(7), 443-9  
CODEN: HKZAA2  
DOCUMENT TYPE: Journal  
LANGUAGE: Japanese  
TI **Microbial production of amino acids**  
. VI. Formation of L-amino acids from  
DL-.alpha.-hydroxycarboxylic acids  
AB Formation of L-amino acids from DL-.alpha.-  
hydroxycarboxylic acids was studied. L-.alpha.-aminobutyric acid  
[1492-24-6] was formed in a medium contg. DL-.alpha.-hydroxybutyric acid  
[600-15-7] by various bacteria belonging to Aerobacter, Bacillus,  
**Corynebacterium**, Escherichia, Flavobacterium, Micrococcus,  
Proteus, Pseudomonas, Sarcina, Staphylococcus, and other genera. A.  
cloacae IAM 1221 was cultured in a medium contg. DL-.alpha.-bromobutyric  
acid [2385-70-8] (hydrolyzed to hydroxybutyric acid).  
L-.alpha.-aminobutyric acid was isolated from the culture broth and

identified by thin-layer chromatog., elementary anal., and by its specific rotation and IR spectrum. Formation of valine [72-18-4], leucine [61-90-5], or phenylalanine [63-91-2] from DL-.alpha.-hydroxycarboxylic acids by *Brevibacterium roseum* ATCC 13825 was studied. Yields (mole) from the cultures were 84.22, 95.7, and 47.7%, resp. An amino-group donor (glutamic acid) was needed besides the bacterial cells and DL-.alpha.-hydroxycarboxylic acid for the enzymic formation of amino acids.

L14 ANSWER 15 OF 26 CAPLUS COPYRIGHT 2001 ACS  
ACCESSION NUMBER: 1975:137747 CAPLUS  
DOCUMENT NUMBER: 82:137747  
TITLE: **Microbial production of amino acids**  
INVENTOR(S): Kubota, Koji; Yoshihara, Yasuhiko; Okada, Hiroshi  
PATENT ASSIGNEE(S): Ajinomoto Co., Inc.  
SOURCE: Japan. Kokai, 5 pp.  
CODEN: JKXXAF  
DOCUMENT TYPE: Patent  
LANGUAGE: Japanese  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 49109585	A2	19741018	JP 1973-24049	19730228
JP 51038796	B4	19761023		

TI **Microbial production of amino acids**

AB **Amino acids** were produced by a microbe cultured in a propionic acid medium. Thus, *Brevibacterium flavum* ATCC 14,067, *Micrococcus glutamicus* ATCC 13,032, **Corynebacterium acetoacidophilum** ATCC 13,870, *Microbacterium ammoniaphilum* ATCC 15,354, and *B. flavum* FERM-P 1684 were cultured with shaking at 31.degree. for 48 hr in a medium (pH 7.5) contg. propionic acid 2, (NH4)2SO4 1, KH2PO4 0.1, MgSO4.cntdot.7H2O 0.04, NaCl 0.1, and soybean protein hydrolysate (total

N = 7%) 0.2% plus biotin 2 and thiamine.cntdot.HCl 200 .mu.g/l. Prodn. of L-glutamic acid by each organism was 4.3, 4.2, 3.9, 4.0, and 2.5 mg/ml, resp. *B. flavum* FERM-P 1684 also produced N-acetylglutamine at 0.4 mg/ml.

IT **Corynebacterium acetoacidophilum**  
(glutamic acid manuf. by, from propionic acid)

L14 ANSWER 16 OF 26 CAPLUS COPYRIGHT 2001 ACS  
ACCESSION NUMBER: 1973:56392 CAPLUS  
DOCUMENT NUMBER: 78:56392  
TITLE: **Microbial production of amino acids** from aromatic compounds.  
I. Screening of aromatic compound-assimilating bacteria  
AUTHOR(S): Yamamoto, Masao; Nishida, Hiroshi; Inui, Taiji;  
Ozaki, Asaichiro  
CORPORATE SOURCE: Cent. Res. Lab., Sanraku-Ocean Co., Ltd., Fujisawa,  
Japan  
SOURCE: Hakko Kogaku Zasshi (1972), 50(12), 868-75

CODEN: HKZAA2

DOCUMENT TYPE: Journal  
LANGUAGE: English

TI **Microbial production of amino acids**  
from aromatic compounds. I. Screening of aromatic compound-assimilating bacteria

AB In an attempt to produce **amino acids** from aromatic compds. by fermn., bacterial stock cultures in this lab. were exmd. for their assimilability of benzoate and salicylate; 96 strains from 97 glutamate-producing cultures assimilated benzoic acid. Then, 10 type-strains of the glutamate-producing strains were tested for their assimilability of 40 aromatic compds. 16 of the compds. were assimilated. These were benzoic acid, m-hydroxybenzoic acid, p-hydroxybenzoic acid, 2,4-dihydroxybenzoic acid, 2,5-dihydroxybenzoic acid, 3,4-dihydroxybenzoic acid, 3,5-dihydroxybenzoic acid, phenylacetic acid, phenylpyruvic acid, .beta.-phenylpropionic acid, cinnamic acid, benzaldehyde, benzyl alc., phenol, catechol, and resorcinol. A sizable amt. of L-glutamic acid

was produced from the assimilated compds. by these glutamate-producing bacteria, benzoate, esp., serving as the best substrate.

IT *Brevibacterium*  
*Brevibacterium lactofermentum*  
**Corynebacterium acetoglutamicum**  
*Microbacterium ammoniaphilum*  
*Micrococcus glutamicus*  
(glutamic acid formation by, from arom. compds.)

L14 ANSWER 17 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1971:84222 CAPLUS

DOCUMENT NUMBER: 74:84222

TITLE: Utilization of hydrocarbons by microorganisms. XXI.  
Biochemical studies of **microbial production** of .alpha.-ketoglutarate,

L-glutamate, and DL-alanine from hydrocarbons

AUTHOR(S): Imada, Yukio; Yamada, Koichi

CORPORATE SOURCE: Fac. Agric., Univ. Tokyo, Tokyo, Japan

SOURCE: Agr. Biol. Chem. (1971), 35(1), 18-26

CODEN: ABCHA6

DOCUMENT TYPE: Journal

LANGUAGE: English

TI Utilization of hydrocarbons by microorganisms. XXI. Biochemical studies of **microbial production** of .alpha.-ketoglutarate, L-glutamate, and DL-alanine from hydrocarbons

AB Strain S10B1 of **Corynebacterium hydrocarbolastus** produced .alpha.-ketoglutaric acid (I), L-Glutamate, and DLAlanine from nAlkanes in a thiam (II) Limited medium supplemented with Fe2+. The replacement of hydrocarbon substrate by sugars such as glucose not only decreased the yields, but also reversed the order of the yields among the 3 products. This phenomenon was explained by a metabolic pathway in relation to the role of II. Slow O uptake in the presence of pyruvate and I by IIDeficient cells supported the presumption that II limitation resulted

in deficiency of a cofactor in the enzymic oxidn. of pyruvate and I. Activities of terminal enzymes in the synthesis of LGlutamate and DLLanine

were detd. and discussed. Three intermediates were detected in the culture broth.

ST **Corynebacterium** ketoglutarate prodn; ketoglutarate prodn

IT    **Corynebacterium**; glutamate prodn **Corynebacterium**; alanine prodn **Corynebacterium**; thiamine **Corynebacterium**; hydrocarbons ultilization bacteria; bacteria hydrocarbons utilization  
IT    **Corynebacterium**  
      (hydrocarboclastus, **amino acids** formation by, from hydrocarbons)  
IT    59-43-8, biological studies  
RL: BIOL (Biological study)  
      (**amino acids** formation from paraffins by **Corynebacterium** hydrocarboclastus in response to)

L14 ANSWER 18 OF 26 CAPLUS COPYRIGHT 2001 ACS  
ACCESSION NUMBER: 1970:475660 CAPLUS  
DOCUMENT NUMBER: 73:75660  
TITLE: **Microbial production of L-glutamic acid**  
PATENT ASSIGNEE(S): Asahi Chemical Industry Co., Ltd.  
SOURCE: Fr. Demande, 11 pp.  
CODEN: FRXXBL  
DOCUMENT TYPE: Patent  
LANGUAGE: French  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
FR 2009795		19700206		

PRIORITY APPLN. INFO.: JP 19680531  
TI **Microbial production of L-glutamic acid**  
AB L-Glutamic acid (I) is prepd. by aerobic cultivation of **Corynebacterium** or *Brevibacterium* in liq. media contg. C1-3 alcs. as C source and penicillin. Thus, *B. vitalumen* var *propanolophilum* ATCC 21391 was grown in a medium contg. PrOH 50, corn steep liquor 4, KH<sub>2</sub>PO<sub>4</sub> 2,  
MgSO<sub>4</sub>.7H<sub>2</sub>O 0.5, Fe<sup>2+</sup> 0.01, Mn<sup>2+</sup> 0.01, urea 4 g/l., with the addn. of 100 .mu.g biotin and penicillin G (K salt) 10 units/l., at 32.degree. and pH 6.5-8.0 with shaking for 96 hr to give 23.1 g I/l. (46.2% based on PrOH). PrOH and penicillin were added in portions during the fermentation. Without penicillin addn., the yield was 6.4% I.  
ST *Brevibacterium glutamate* prodn; *glutamate* prodn *Brevibacterium*; **amino acids** **Corynebacterium**; **Corynebacterium amino acids**; penicillin bacteria glutamate  
IT **Corynebacterium**  
      (melassecola and petrophylum, glutamic acid manuf. by)

L14 ANSWER 19 OF 26 CAPLUS COPYRIGHT 2001 ACS  
ACCESSION NUMBER: 1970:508239 CAPLUS  
DOCUMENT NUMBER: 73:108239  
TITLE: **Microbial production of L-threonine**  
INVENTOR(S): Nakayama, Kiyoshi; Kase, Hiroshi  
PATENT ASSIGNEE(S): Kyowa Fermentation Industry Co. Ltd.  
SOURCE: Ger. Offen., 22 pp.  
CODEN: GWXXBX  
DOCUMENT TYPE: Patent  
LANGUAGE: German  
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
DE 1817666	A	19700827	DE 1968-1817666	19681224
TI	<b>Microbial production</b> of L-threonine			
AB	Various microorganisms, e.g. Aerobacter [Enterobacter] aerogenes, <i>Serratia</i> marcescens, or Arthrobacter paraffineus, cultured for producing L-threonine required 2 or 3 of the <b>amino acids</b> isoleucine, methionine, lysine, or diaminopimelic acid. The microorganisms were cultured aerobically in an aq. medium contg. the optimal (or less) amts. of the required <b>amino acids</b> . Thus, <i>E. aerogenes</i> NM-IS-5 (ATCC 21,215) was cultured 96 hr at 30.degree. in medium contg. glucose 5, (NH4)2SO4 1.4, KH2PO4 0.05, K2HPO4 0.05, MgSO4·7H2O 0.025, FeSO4·7H2O 0.001, MnSO4·4H2O 0.001, and CaCO3 2% and isoleucine 50, methionine 100, and diaminopimelic acid 200 mg/l. to give 7.8 g L-threonine/l.			
ST	<b>microbial prodn</b> threonine; threonine <b>microbial</b> prodn; Aerobacter threonine fermn; amino acid prodn fermn			
IT	<b>Corynebacterium</b> (glutamicum, threonine manuf. by)			
L14	ANSWER 20 OF 26 CAPLUS COPYRIGHT 2001 ACS			
ACCESSION NUMBER:	1970:401150 CAPLUS			
DOCUMENT NUMBER:	73:1150			
TITLE:	<b>Microbial production</b> of L-threonine. II. Production by .alpha.-amino-.beta.-			
	hydroxyvaleric acid resistant mutants of glutamate producing bacteria			
AUTHOR(S):	Shio, Isamu; Nakamori, Shigeru			
CORPORATE SOURCE:	Cent. Res. Lab., Ajinomoto Co., Inc., Kawasaki, Japan			
SOURCE:	Agr. Biol. Chem. (1970), 34(3), 448-56			
DOCUMENT TYPE:	CODEN: ABCHA6			
LANGUAGE:	Journal			
TI	<b>Microbial production</b> of L-threonine. II. Production by .alpha.-amino-.beta.-hydroxyvaleric acid resistant mutants of glutamate producing bacteria			
AB	A mutant strain of <i>Brevibacterium flavum</i> was able to grow in a medium contg. 5 mg DL-threo-.alpha.-amino-.beta.-hydroxyvaleric acid (AHV)/ml; 1 mg AHV/ml inhibited the growth of the parental strain by >90%. Further treatment of the AHV-resistant strain with the mutagen, N-methyl-N'-nitro-N-nitrosoguanidine, produced a bacterial strain that was able to grow on 8 mg AHV/ml; this mutant produced 13.5 g L-threonine/l., an amt. 30% more than that produced by the parental strain. A similarly derived mutant of <b>Corynebacterium acetoacidophilum</b> resistant to AHV produced 6.1 g threonine/l. Other <b>amino acids</b> biosynthesized by the bacteria were discussed in relation to the regulation of threonine synthesis.			
ST	threonine prodn bacterial; <b>corynebacterium</b> threonine prodn; <i>Brevibacterium</i> threonine prodn; mutations bacteria threonine; bacteria mutations threonine; aminohydroxyvalerate bacteria			
IT	<b>Corynebacterium</b> (acetoacidophilum, tryptophan formation from aminohydroxyvaleric acid			

by)

L14 ANSWER 21 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS |  
ACCESSION NUMBER: 1970:123860 BIOSIS  
DOCUMENT NUMBER: BA51:33860  
TITLE: **MICROBIAL PRODUCTION OF AMINO**  
-ACIDS FROM HYDRO CARBONS PART 4 L GLUTAMIC-ACID  
PRODUCTION BY **CORYNEBACTERIUM-HYDROCARBOCLASTUS-R-**  
7.  
AUTHOR(S): SHII O I; UCHIO R  
SOURCE: AMINO ACID NUCLEIC ACID, (1969) (19), 88-96.  
CODEN: HATAA4. ISSN: 0517-6174.  
FILE SEGMENT: BA; OLD  
LANGUAGE: Unavailable  
TI **MICROBIAL PRODUCTION OF AMINO-ACIDS**  
FROM HYDRO CARBONS PART 4 L GLUTAMIC-ACID PRODUCTION BY  
**CORYNEBACTERIUM-HYDROCARBOCLASTUS-R-7.**

L14 ANSWER 22 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS DUPLICATE 2  
ACCESSION NUMBER: 1970:106213 BIOSIS  
DOCUMENT NUMBER: BA51:16213  
TITLE: **MICROBIAL PRODUCTION OF AMINO**  
-ACIDS FROM HYDRO CARBONS PART 4 L GLUTAMIC-ACID  
PRODUCTION BY **CORYNEBACTERIUM-HYDROCARBOCLASTUS**  
R-7.  
AUTHOR(S): SHII O I; UCHIO R  
SOURCE: J GEN APPL MICROBIOL, (1969) 15 (1), 65-84.  
CODEN: JGAMA9. ISSN: 0022-1260.  
FILE SEGMENT: BA; OLD  
LANGUAGE: Unavailable  
TI **MICROBIAL PRODUCTION OF AMINO-ACIDS**  
FROM HYDRO CARBONS PART 4 L GLUTAMIC-ACID PRODUCTION BY  
**CORYNEBACTERIUM-HYDROCARBOCLASTUS R-7.**

L14 ANSWER 23 OF 26 CAPLUS COPYRIGHT 2001 ACS  
ACCESSION NUMBER: 1967:514494 CAPLUS  
DOCUMENT NUMBER: 67:114494  
TITLE: **Microbial production of**  
**amino acids from hydrocarbons. III.**  
L-Ornithine production by an arginine auxotrophic  
mutant of **Corynebacterium hydrocarboclastus**  
Ishu, Ryosuke; Ishii, Ryosuke; Shio, Isamu  
Ajinomoto Co., Inc., Kawasaki, Japan  
J. Gen. Appl. Microbiol. (1967), 13(3), 3303-12  
CODEN: JGAMA9  
DOCUMENT TYPE: Journal  
LANGUAGE: English  
TI **Microbial production of amino acids**  
from hydrocarbons. III. L-Ornithine production by an arginine  
auxotrophic mutant of **Corynebacterium hydrocarboclastus**  
AB cf. CA 67: 89718u. The arginine auxotrophic mutant strain RN-362 of C.  
hydrocarboclastus R-7 was used to study L-ornithine production from  
hydrocarbons, in a fermentation medium contg. various n-alkanes.  
L-Ornithine production required L-arginine at the optimum level of  
0.5-1.0  
g./l. of medium; an excess inhibited the biosynthesis of L-ornithine.  
(NH4)2HPO4 was the best source of N and, at 2% in a neutral to slightly  
acidic pH, gave the highest level of L-ornithine production and cell

growth; NH<sub>4</sub>OAc, KNO<sub>3</sub>, and (NH<sub>4</sub>)<sub>2</sub>CO<sub>3</sub> proved less suitable because of a drop in pH along with the accumulation of a large amt. of .alpha.-ketoglutaric acid, pyruvic acid, and proline in the growth medium. Of 17 C sources, n-tetradecane best supported cell growth and L-ornithine production and the other C<sub>13</sub>-C<sub>17</sub> n-alkanes did so moderately, while kerosene and light oil produced good cell growth but only a small amt. of L-ornithine.

Addn. of 3 g. yeast ext. and 0.5 g. L-arginine-HCl to 1 l. of medium enhanced L-ornithine production. A similar effect was achieved by replacing the yeast ext. with various **amino acids** at 0.01% in the medium. L-Methionine was most effective for the production of L-ornithine, while L-lysine, L-cysteine, L-cystine, L-histidine, and L-phenylalanine were less so, in decreasing order. **Amino acids** enhance L-ornithine production by stimulating hydrocarbon oxidn. and cell growth.

ST HYDROCARBONS USE BACTERIA; BACTERIA HYDROCARBONS USE; ALKANES USE BACTERIA; **AMINO ACIDS** PRODN HYDROCARBONS; ORNITHINE PRODN HYDROCARBONS; PARAFFINS UTILIZATION BACTERIA

IT **Corynebacterium**  
(hydrocarboclastus, ornithine formation from hydrocarbons by)

IT Hydrocarbons, biological studies

RL: BIOL (Biological study)  
(ornithine formation from, by **Corynebacterium**  
hydrocarboclastus)

IT 70-26-8

RL: FORM (Formation, nonpreparative)  
(formation of, from hydrocarbons by **Corynebacterium**  
hydrocarboclastus)

L14 ANSWER 24 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1967:489718 CAPLUS

DOCUMENT NUMBER: 67:89718

TITLE: **Microbial production of amino acids from hydrocarbons. II.**  
Isolationf good hycarbon utilizers and amino acid production by their auxotrophs

AUTHOR(S): Ishii, Ryosuke; Otsuka, Shinichiro; Shio, Isamu

CORPORATE SOURCE: Central Res. Labs., Ajinomoto Co., Inc., Kawasaki,  
Japan

SOURCE: J. Gen. Appl. Microbiol. (1967), 13(2), 217-25  
CODEN: JGAMA9

DOCUMENT TYPE: Journal

LANGUAGE: English

TI **Microbial production of amino acids**  
from hydrocarbons. II. Isolationf good hycarbon utilizers and amino acid production by their auxotrophs

AB cf. CA 59: 14313h. Nine microorganisms, which showed good growth on long-chain aliphatic hydrocarbons, were isolated by an enrichment culture method, followed by a single colony isolation technique. They included 5 strains of Alcaligenes marshallii, 2 strains of **Corynebacterium** hydrocarboclastus, and 2 strains of yeast. Various auxotrophic mutants were derived from these microorganisms. The mutants accumulated the following **amino acids** from aliphatic hydrocarbons; L-ornithine, L-valine, L-glutamic acid, L-leucine, L-tyrosine, L-alanine, L-proline, L-aspartic acid, and L-lysine.

ST BACTERIA AMINO ACID PRODN; AMINO ACID PRODN BACTERIA; HYDROCARBONS

IT      **AMINO ACIDS; ALIPHATICS BACTERIA METAB**  
IT      **Corynebacterium**  
          (hydrocarboblastus, amino acid fermentation of hydrocarbons by)  
IT      **Amino acids, preparation**  
RL: BMF (Bioindustrial manufacture); BIOL (Biological study); PREP  
(Preparation)  
      (manuf. of, by fermentation of hydrocarbons)

L14 ANSWER 25 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:            1966:22870 CAPLUS  
DOCUMENT NUMBER:            64:22870  
ORIGINAL REFERENCE NO.:    64:4230g-h,4231a  
TITLE:                        **Microbial production of**  
                              nucleotides  
INVENTOR(S):                Masuo, Eitaro; Okabayashi, Tadashi  
PATENT ASSIGNEE(S):        Shionogi & Co., Ltd.  
SOURCE:                     10 pp.  
DOCUMENT TYPE:             Patent  
LANGUAGE:                    Unavailable  
FAMILY ACC. NUM. COUNT:    1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 40010957		19650601	JP	19591214

TI      **Microbial production** of nucleotides  
AB      Some bacteria strains of high nucleotide-forming activity were detected based on the results of the test developed by the authors, and compns. of media for promoting accumulation of nucleotides were also investigated. To evaluate the nucleotide-forming activity of bacteria, cells of nonexacting purine (I) auxotrophic mutant B 96 of Escherichia coli were mixed into the synthetic medium contg. no I for testing strains. The activity of nucleotide accumulation of the strains increased as the growth of the mutant increased. By this procedure, the following strains were found to be suitable for nucleotide production: Bacillus subtilis IFO 3061, B. firmus IFO 3330, B. circulans IFO 3342, B. megaterium IFO 3003, Alcaligenes viscosus AN-14, A. metalcaligenes 1021, Serratia marcescens 1008, S. plymuthica IFO 3055, Bacterium ketoglutaricum 1041, and new species of Brevibacterium and **Corynebacterium**. For promoting nucleotide production with these strains, **amino acids**, esp. L-glutamic acid (II), are necessary in the medium. Proteins or peptides contg. II are also effective for the strains having sufficient protease. Sufficient content of PO43- at pH 5.0-7.5 is also necessary for the medium. By cultivation under these conditions, AMP, CDP, UMP, and UDP are obtained.

L14 ANSWER 26 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:            1963:476777 CAPLUS  
DOCUMENT NUMBER:            59:76777  
ORIGINAL REFERENCE NO.:    59:14313h,14314a  
TITLE:                        **Microbial production of**  
                              **amino acids** from hydrocarbons. I.  
                              Preliminary screening of glutamic acid-producing  
                              bacteria  
AUTHOR(S):                  Shioi, Isamu; Otsuka, Shinichiro; Ishii, Ryosuke;

CORPORATE SOURCE: Katsuya, Nobu; Iizuka, Hiroshi  
Ajinomoto Co., Inc., Kawasaki, Japan  
SOURCE: J. Gen. Appl. Microbiol. (Tokyo) (1963), 9, 23-30  
DOCUMENT TYPE: Journal  
LANGUAGE: Unavailable  
TI **Microbial production of amino acids**  
from hydrocarbons. I. Preliminary screening of glutamic acid-producing  
bacteria  
AB Various bacteria utilized kerosene, light oil, heavy oil, and liquid  
paraffin as the only C source for growth and formation of L-glutamic acid  
(I). The highest level of I (281 .gamma./ml.) was obtained from kerosene  
by a strain of **Corynebacterium** hydrocarboclastus.

=> dis L15 1-13 ibib kwic

L15 ANSWER 1 OF 13 BIOSIS COPYRIGHT 2001 BIOSIS  
ACCESSION NUMBER: 2001:314459 BIOSIS  
DOCUMENT NUMBER: PREV200100314459  
TITLE: Effect of gluconic acid as a secondary carbon source on  
non-growing L-**lysine** producers cells of  
**Corynebacterium** glutamicum. Purification and  
properties of 6-phosphogluconate dehydrogenase.  
AUTHOR(S): Bianchi, Daniella; Bertrand, Olivier; Haupt, Karsten;  
Coello, Nereida (1)  
CORPORATE SOURCE: (1) Instituto de Biologia Experimental, Universidad  
Central  
de Venezuela, Caracas, 1041-A: ncoello@uole.com.ve  
Venezuela  
SOURCE: Enzyme and Microbial Technology, (June 7, 2001) Vol. 28,  
No. 9-10, pp. 754-759. print.  
ISSN: 0141-0229.  
DOCUMENT TYPE: Article  
LANGUAGE: English  
SUMMARY LANGUAGE: English  
TI Effect of gluconic acid as a secondary carbon source on non-growing L-  
**lysine** producers cells of **Corynebacterium** glutamicum.  
Purification and properties of 6-phosphogluconate dehydrogenase.  
AB We studied the production of L-**lysine** in **Corynebacterium**  
glutamicum ATCC 21543 non growing cells obtained by nutrient limitation.  
Statistical analysis revealed significant differences in the L-  
**lysine** titers of glucose, gluconic acid or glucose-gluconic acid  
cultures. Higher L-**lysine** titer obtained in batch cultures with  
mixed carbon sources or gluconic acid alone were found to be associated  
with a . . . dehydrogenase activity (6PGDH, E.C.1.1.1.44). This enzyme  
is a pivotal enzyme within the hexose monophosphate pathway, and thus of  
importance for L-**lysine** production. 6PGDH was purified and  
characterized. The purified enzyme migrates as a single band on sodium  
dodecyl sulfate-polyacrylamide gel electrophoresis. . . .  
IT . . .  
Bioprocess Engineering; Methods and Techniques; Nutrition  
IT Chemicals & Biochemicals  
6-phosphogluconate dehydrogenase: amino acid sequence, analysis,  
molecular properties, pH, purification; L-**lysine**:  
**microbial production**, yield; **amino acids**: analysis; carbon sources; gluconic acid: secondary  
carbon source  
ORGN . . .

Microorganisms; Irregular Nonsporing Gram-Positive Rods: Actinomycetes and Related Organisms, Eubacteria, Bacteria, Microorganisms; Microorganisms

ORGN Organism Name

Bacillus subtilis (Endospore-forming Gram-Positives);

**Corynebacterium** glutamicum (Irregular Nonsporing Gram-Positive Rods): non-growing cells; Escherichia coli (Enterobacteriaceae); bacteria (Bacteria); microorganisms (Microorganisms)

ORGN Organism Superterms

Bacteria; Eubacteria; Microorganisms

RN 9001-82-5Q (6-PHOSPHOGLUCONATE DEHYDROGENASE)

9073-95-4Q (6-PHOSPHOGLUCONATE DEHYDROGENASE)

56-87-1 (L-**LYSINE**)

526-95-4 (GLUCONIC ACID)

L15 ANSWER 2 OF 13 BIOSIS COPYRIGHT 2001 BIOSIS

ACCESSION NUMBER: 2001:174925 BIOSIS

DOCUMENT NUMBER: PREV200100174925

TITLE: MALDI-TOF MS for quantification of substrates and products in cultivations of **Corynebacterium** glutamicum.

AUTHOR(S): Wittmann, Christoph (1); Heinze, Elmar

CORPORATE SOURCE: (1) Biochemical Engineering Institute, Saarland

University,

66041, Saarbruecken: c.wittmann@rz.uni-sb.de Germany

SOURCE: Biotechnology and Bioengineering, (March 20, 2001) Vol.

72,

No. 6, pp. 642-647. print.

ISSN: 0006-3592.

DOCUMENT TYPE: Article

LANGUAGE: English

SUMMARY LANGUAGE: English

TI MALDI-TOF MS for quantification of substrates and products in cultivations

of **Corynebacterium** glutamicum.

AB The application of MALDI-TOF MS for the quantification of **lysine**, alanine, and glucose is described. The method is based on using stable isotopes as internal standards and allows fast, sensitive, . . . concentrations of the analytes between 10 muM and 100 mM. The mean standard deviations from five replicates each were 4.3% (**lysine**), 3.7% (alanine), and 3.2% (glucose). In addition, sucrose could be measured by MALDI-TOF MS, but was not quantified due to. . .

IT Major Concepts

Biochemistry and Molecular Biophysics; Bioprocess Engineering; Methods and Techniques

IT Chemicals & Biochemicals

**amino acids: microbial production**

, quantitative analysis; products: quantitative analysis; substrates: quantitative analysis

L15 ANSWER 3 OF 13 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:761605 CAPLUS

DOCUMENT NUMBER: 134:99608

TITLE: Development and use of miniaturized parallel experiment technology for bioprocess development

AUTHOR(S): Altenbach-Rehm, Jutta

CORPORATE SOURCE: Institut fur Biotechnologie, Julich, JUL-3782, Germany

SOURCE: Ber. Forschungszent. Juelich (2000), Juel-3782, a-f,

i-iv, 1-233  
CODEN: FJBEE5; ISSN: 0366-0885

DOCUMENT TYPE:

Report

LANGUAGE:

German

AB The fed-batch technique is nowadays the std. operation mode for high performance **microbial prodn.** processes. Shake flasks are widely used a simple bioreactors in batch process development. Because of uncontrolled changes in pH and reduced oxygen transfer rates parallel operated shake flasks can usually not be applied for microbial fed-batch process development. Due to the need to operate controlled stirred tank reactors the development of specific fermn. strategies is expensive and time consuming. To address these issues a miniature fed-batch technique was developed in cooperation with DASGIP mbH, Julich, and INFORS AG, Basel. Controlled batch and fed-batch fermns. can be performed in 16 parallel small scale bioreactors. The new technique allows the feeding of up to 4 different substrates and parallel pH control

based on user-defined profiles. The feeding assembly is sterilized chem. using dimethylcarbonate. To overcome the limitations of shake flasks, small scale bubble columns were developed. Gas distribution is performed with a new type of sterile sparger. Transferring fed-batch fermns. from small scale bubble columns to a stirred tank reactor a scale up factor of 20 was achieved. *Escherichia coli* K12 was chosen to test the new parallel

bioreactor technique. Compared to shake flask fermns. the cell concn.

was

50% higher due to efficient oxygen transfer. In fed-batch fermns. with pH-controlled substrate feeding up to 35 g/l DCW were achieved. For the 1st time, a parallel optimization of feeding profiles in parallel small scale fed-batch expts. with successful scale-up to a lab. bioreactor was performed. *Escherichia coli* BL 21 (DE3) pLySS produces the recombinant enzyme GDP-.alpha.-D-mannose-pyrophosphorylase after induction with isopropyl-.beta.-D-thiogalactoside (IPTG). Single induction with 0,5 mM IPTG resulted in a low specific enzyme activity of 1,6 U/g DCW (dry cell wt.). For the optimization of enzyme expression a genetic algorithm was used. A final enzyme activity of 111 U/g DCW was achieved for optimal substrate and inducer feeding profiles. To demonstrate the advantages of this new parallel bioreactor technique different strains of industrial relevance were investigated. **Corynebacterium glutamicum**, *Staphylococcus carnosus* and *Ashbya gossypii*.

ST bioreactor bubble column fed batch fermn; GDP mannose pyrophosphorylase bubble column fed batch; *Staphylococcus* bubble column fed batch fermn; isoleucine bubble column fed batch **Corynebacterium**; riboflavin bubble column fed batch *Ashbya*

IT **Amino acids**, biological studies

RL: BPR (Biological process); BIOL (Biological study); PROC (Process)  
(amino acid consumption in riboflavin prodn. by *Ashbya gossypii* in parallel bubble columns with fed-batch technique)

IT **Corynebacterium glutamicum**

(L-isoleucine prodn. by **Corynebacterium glutamicum** in parallel bubble columns with fed-batch technique)

IT 56-40-6, Glycine, biological studies 56-41-7, Alanine, biological studies 56-45-1, Serine, biological studies 56-84-8, Aspartic acid, biological studies 56-85-9, Glutamine, biological studies 56-86-0, Glutamic acid, biological studies 56-87-1, **Lysine**, biological studies 60-18-4, Tyrosine, biological studies 63-68-3, Methionine, biological studies 63-91-2, Phenylalanine, biological studies

72-18-4,

Valine, biological studies 72-19-5, Threonine, biological studies  
 73-22-3, Tryptophane, biological studies  
 RL: BPR (Biological process); BIOL (Biological study); PROC (Process)  
     (amino acid consumption in riboflavin prodn. by *Ashybya gossypii* in  
     parallel bubble columns with fed-batch technique)

IT 73-32-5P, L-Isoleucine, biological studies  
 RL: BMF (Bioindustrial manufacture); BPR (Biological process); BIOL  
     (Biological study); PREP (Preparation); PROC (Process)  
     (L-isoleucine prodn. by *Corynebacterium glutamicum* in  
     parallel bubble columns with fed-batch technique, amino acid  
     consumption in riboflavin prodn. by *Ashybya gossypii*)

IT 61-90-5, L-Leucine, biological studies  
 RL: BPR (Biological process); MFM (Metabolic formation); BIOL (Biological  
     study); FORM (Formation, nonpreparative); PROC (Process)  
     (L-isoleucine prodn. by *Corynebacterium glutamicum* in  
     parallel bubble columns with fed-batch technique, amino acid  
     consumption in riboflavin prodn. by *Ashybya gossypii*)

L15 ANSWER 4 OF 13 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1999:244776 CAPLUS  
 DOCUMENT NUMBER: 130:266420  
 TITLE: Method for **microbial production of amino acids** of the aspartate and/or glutamate family and agents which can be used in said method  
 INVENTOR(S): Eikmanns, Bernd; Peters-Wendisch, Petra; Sahm, Hermann  
 PATENT ASSIGNEE(S): Forschungszentrum Julich G.m.b.H., Germany  
 SOURCE: PCT Int. Appl., 40 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: German  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9918228	A2	19990415	WO 1998-EP6210	19980930
WO 9918228	A3	19990520		
W:	AU, BR, CA, CN, HU, ID, JP, KR, MX, RU, SK, US			
RW:	AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE			
DE 19831609	A1	19990415	DE 1998-19831609	19980714
AU 9911482	A1	19990427	AU 1999-11482	19980930
EP 1015621	A2	20000705	EP 1998-954301	19980930
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI			
BR 9813021	A	20000815	BR 1998-13021	19980930
PRIORITY APPLN. INFO.:			DE 1997-19743894 A	19971004
			DE 1998-19831609 A	19980714
			WO 1998-EP6210 W	19980930

TI Method for **microbial production of amino acids** of the aspartate and/or glutamate family and agents which can be used in said method  
 AB The invention relates to a method for **microbial prodn. of amino acids** of the aspartate and/or glutamate family in which the pyruvate carboxylase activity is increased by genetically changing the enzyme and/or the pyruvate carboxylase gene

expression of a microorganism which produces the corresponding amino acid.

In addn., the invention relates to a pyruvate carboxylase gene and addnl. agents which can be used in the inventive method.

ST amino acid fermn **Corynebacterium** pyruvate carboxylase genetic engineering

IT **Corynebacterium** glutamicum  
Fermentation  
(**microbial prodn. of amino acids**  
of the aspartate and/or glutamate family and agents which can be used in said method)

IT Amino acids, preparation  
RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(**microbial prodn. of amino acids**  
of the aspartate and/or glutamate family and agents which can be used in said method)

IT Genetic engineering  
(**microbial prodn. of amino acids**  
of the aspartate and/or glutamate family and modification of **Corynebacterium** pyc gene in said method)

IT Genes (microbial)  
RL: BPR (Biological process); BIOL (Biological study); PROC (Process)  
(pyc; **microbial prodn. of amino acids** of the aspartate and/or glutamate family and modification of **Corynebacterium** pyc gene in said method)

IT 56-86-0P, L-Glutamic acid, preparation 56-87-1P, L-**Lysine**, preparation 72-19-5P, L-Threonine, preparation 74-79-3P, L-Arginine, preparation 672-15-1P, L-Homoserine  
RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(**microbial prodn. of amino acids**  
of the aspartate and/or glutamate family and agents which can be used in said method)

IT 9014-19-1, Pyruvate carboxylase 204116-18-7 208541-81-5  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(**microbial prodn. of amino acids**  
of the aspartate and/or glutamate family and agents which can be used in said method)

L15 ANSWER 5 OF 13 BIOSIS COPYRIGHT 2001 BIOSIS  
ACCESSION NUMBER: 1999:175414 BIOSIS  
DOCUMENT NUMBER: PREV199900175414  
TITLE: Cloning of the transketolase gene and the effect of its dosage on aromatic amino acid production in **Corynebacterium** glutamicum.  
AUTHOR(S): Ikeda, M. (1); Okamoto, K.; Katsumata, R.  
CORPORATE SOURCE: (1) Technical Research Laboratories, Kyowa Hakko Kogyo Co., Ltd., Hofu, Yamaguchi, 747-8522 Japan  
SOURCE: Applied Microbiology and Biotechnology, (Feb., 1999) Vol. 51, No. 2, pp. 201-206.  
ISSN: 0175-7598.  
DOCUMENT TYPE: Article  
LANGUAGE: English  
TI Cloning of the transketolase gene and the effect of its dosage on aromatic amino acid production in **Corynebacterium** glutamicum.

AB. . . enzyme of the non-oxidative pentose phosphate pathway. The effect of its overexpression on aromatic amino acid production was investigated in **Corynebacterium glutamicum**, a typical amino-acid-producing organism. For this purpose, the transketolase gene of the organism was cloned on the basis of. . . as a protein of approximately 83kDa in proportion to the copy numbers. Introduction of the plasmids into a tryptophan and **lysine** co-producer resulted in copy-dependent increases in tryptophan production along with concomitant decreases in **lysine** production. Furthermore, the presence of the gene in high copy numbers enabled tyrosine, phenylalanine and tryptophan producers to accumulate 5%-20% more aromatic **amino acids**. These results indicate that overexpressed transketolase activity operates to redirect the glycolytic intermediates toward the nonoxidative pentose phosphate pathway in. . .

IT Major Concepts

Bioprocess Engineering; Enzymology (Biochemistry and Molecular Biophysics); Molecular Genetics (Biochemistry and Molecular Biophysics)

IT Chemicals & Biochemicals

aromatic **amino acids**: **microbial production**; transketolase [EC 2.2.1.1]; **Corynebacterium transketolase gene** (Irregular Nonsporing Gram-Positive Rods)

L15 ANSWER 6 OF 13 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1995:674124 CAPLUS  
 DOCUMENT NUMBER: 123:54314  
 TITLE: Enhancement of reduced NADP production for enhanced **microbial production** of biochemicals  
 INVENTOR(S): Kojima, Hiroyuki; Totsuka, Kazuhiko  
 PATENT ASSIGNEE(S): Ajinomoto Co., Inc., Japan  
 SOURCE: PCT Int. Appl., 32 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: Japanese  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9511985	A1	19950504	WO 1994-JP1791	19941026
W: AU, BR, CA, CN, CZ, HU, JP, KR, PL, RU, SK, US, VN				
RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
CA 2175042	AA	19950504	CA 1994-2175042	19941026
AU 9480026	A1	19950522	AU 1994-80026	19941026
AU 687458	B2	19980226		
EP 733712	A1	19960925	EP 1994-931158	19941026
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT,				
SE				
BR 9407907	A	19961126	BR 1994-7907	19941026
HU 74840	A2	19970228	HU 1996-1085	19941026
ZA 9503350	A	19961025	ZA 1995-3350	19950425
US 5830716	A	19981103	US 1996-619521	19960429
CN 1139956	A	19970108	CN 1994-194707	19961026
PRIORITY APPLN. INFO.:			JP 1993-270828	19931028
			WO 1994-JP1791	19941026

TI Enhancement of reduced NADP production for enhanced **microbial production** of biochemicals

AB The productivity of such substances as L-amino acids, antibiotics, vitamins, growth factors and physiol. active substances in the fermn. using a microorganism is improved by improving the productivity of reduced NADP in the cells of the microorganisms. Construction of pMW::THY contg. the Escherichia coli transhydrogenase gene, and introduction of the plasmid into the L-threonine-producing Escherichia coli B-3996 were shown. The recombinant E. coli B-3996 produced L-threonine .apprx.10% higher than did the parental strain.

IT **Corynebacterium glutamicum**  
Escherichia coli  
Fermentation  
(enhancement of reduced NADP prodn. for enhanced **microbial prodn. of biochems.**)

IT **Amino acids, preparation**  
RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(enhancement of reduced NADP prodn. for enhanced **microbial prodn. of biochems.**)

IT Plasmid and Episome  
(pHSG::THY; enhancement of reduced NADP prodn. for enhanced **microbial prodn. of biochems.**)

IT Plasmid and Episome  
(pMW::THY; enhancement of reduced NADP prodn. for enhanced **microbial prodn. of biochems.**)

IT Plasmid and Episome  
(pSU::THY; enhancement of reduced NADP prodn. for enhanced **microbial prodn. of biochems.**)

IT 9014-18-0, Nicotinamide nucleotide transhydrogenase  
RL: BAC (Biological activity or effector, except adverse); BIOL (Biological study)  
(enhancement of reduced NADP prodn. for enhanced **microbial prodn. of biochems.**)

IT 56-86-0P, L-Glutamic acid, preparation 56-87-1P, L-Lysine, preparation 61-90-5P, L-Leucine, preparation 63-91-2P, L-Phenylalanine, preparation 72-18-4P, L-Valine, preparation 72-19-5P,  
L-Threonine, preparation 73-32-5P, L-Isoleucine, preparation  
RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(enhancement of reduced NADP prodn. for enhanced **microbial prodn. of biochems.**)

IT 53-59-8P, NADP  
RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(reduced; enhancement of reduced NADP prodn. for enhanced **microbial prodn. of biochems.**)

L15 ANSWER 7 OF 13 SCISEARCH COPYRIGHT 2001 ISI (R)  
ACCESSION NUMBER: 95:184306 SCISEARCH  
THE GENUINE ARTICLE: QK574  
TITLE: METABOLIC DESIGN IN AMINO-ACID PRODUCING BACTERIUM  
**CORYNEBACTERIUM-GLUTAMICUM**  
AUTHOR: SAHM H (Reprint); EGGLING L; EIKMANNS B; KRAMER R  
CORPORATE SOURCE: KFA JULICH GMBH, FORSCHUNGSZENTRUM, INST BIOTECHNOL,  
D-52425 JULICH, GERMANY (Reprint)  
COUNTRY OF AUTHOR: GERMANY  
SOURCE: FEMS MICROBIOLOGY REVIEWS, (FEB 1995) Vol. 16, No. 2-3,

pp. 243-252.  
ISSN: 0168-6445.

DOCUMENT TYPE: Article; Journal  
FILE SEGMENT: LIFE  
LANGUAGE: ENGLISH  
REFERENCE COUNT: 36

\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

TI METABOLIC DESIGN IN AMINO-ACID PRODUCING BACTERIUM **CORYNEBACTERIUM**  
-GLUTAMICUM

AB The Gram-positive bacterium **Corynebacterium glutamicum** is used for the industrial production of **amino acids**, e.g. of L-glutamate and L-**lysine**. In the last 10 years, genetic engineering and amplification of relevant structural genes have become fascinating methods for the construction of strains with desired genotypes. By cloning and expressing the various genes of the L-**lysine** pathway in C. glutamicum we could demonstrate that an increase of the flux of L-aspartate semialdehyde to L-**lysine** could be obtained in strains with increased dehydrodipicolinate synthase activity. By combined overexpression of deregulated aspartate kinase and dihydrodipicolinate synthase, the L-**lysine** secretion could be increased (10-20%). Recently we detected that in C. glutamicum two pathways exist for the synthesis of DL-diaminopimelate and L-**lysine**. Mutants defective in one pathway are still able to synthesize enough L-**lysine** for growth, but the L-**lysine** secretion is reduced to 50-70%. Using NMR spectroscopy, we could calculate

how much of the L-**lysine** secreted into the medium is synthesized via each pathway. Amplification of the feedback inhibition-insensitive homoserine dehydrogenase and homoserine kinase in a high L-**lysine** overproducing strain enabled channelling of the carbon flow from the intermediate aspartate semialdehyde towards homoserine, resulting in a high accumulation. . . acid overproduction, the secretion into the culture medium also has to be noted. Recently it could be demonstrated that L-glutamate, L-**lysine** and L-isoleucine are not secreted via passive diffusion but via specific active carrier systems. Analysis of **lysine**-overproducing C. glutamicum strains indicates that this secretion carrier has a strong influence on the overproduction of this amino acid. Thus, . . .

ST Author Keywords: **CORYNEBACTERIUM GLUTAMICUM**; AMINO ACID PRODUCTION; METABOLIC DESIGN; L-**LYSINE**; L-THREONINE; L-ISOLEUCINE

STP KeyWords Plus (R): L-THREONINE; L-**LYSINE**; BREVIBACTERIUM-LACTOFERMENTUM; HOMOSERINE DEHYDROGENASE; **MICROBIAL** PRODUCTION; RESISTANT MUTANTS; SPLIT PATHWAY; BIOSYNTHESIS; ISOLEUCINE; GENES

L15 ANSWER 8 OF 13 SCISEARCH COPYRIGHT 2001 ISI (R)

ACCESSION NUMBER: 96:7187 SCISEARCH

THE GENUINE ARTICLE: TJ545

TITLE: METABOLIC DESIGN IN THE AMINO-ACID-PRODUCING BACTERIUM **CORYNEBACTERIUM-GLUTAMICUM**

AUTHOR: SAHM H (Reprint)

CORPORATE SOURCE: KFA JULICH GMBH, FORSCHUNGSZENTRUM, INST BIOTECHNOL, D-52425 JULICH, GERMANY (Reprint)

COUNTRY OF AUTHOR: GERMANY

SOURCE: FOLIA MICROBIOLOGICA, (1995) Vol. 40, No. 1, pp. 23-30.  
ISSN: 0015-5632.

DOCUMENT TYPE: Article; Journal

FILE SEGMENT: LIFE; AGRI  
LANGUAGE: ENGLISH  
REFERENCE COUNT: 26  
\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

TI METABOLIC DESIGN IN THE AMINO-ACID-PRODUCING BACTERIUM  
**CORYNEBACTERIUM-GLUTAMICUM**

AB The Gram-positive bacterium **Corynebacterium glutamicum** is used for the industrial production of **amino acids**, e.g. of L-glutamate and L-**lysine**. By cloning and expressing the various genes of the L-**lysine** pathway in *C. glutamicum* we could demonstrate that an increase of the flux of L-4-aspartaldehyde to L-**lysine** could be obtained in strains with increased dihydro-dipicolinate synthase activity. Recently we detected that in *C. glutamicum* two pathways exist for the synthesis of DL-2,6-diaminopimelate and L-**lysine**. Mutants defective in one pathway are still able to synthesize enough L-**lysine** for growth but the L-**lysine** secretion is reduced to 50-70 %. Using NMR-spectroscopy we could calculate how much of the L-**lysine** secreted into the medium is synthesized via the one and the other pathway. Amplification of the feedback-inhibition-insensitive-homoserine dehydrogenase and homoserine kinase in a high L-**lysine**-overproducing strain made it possible to channel the carbon now from the intermediate 4-aspartaldehyde toward homoserine, resulting in a high. . .

STP KeyWords Plus (R): L-THREONINE; BREVIBACTERIUM-LACTOFERMENTUM; HOMOSERINE DEHYDROGENASE; **MICROBIAL PRODUCTION; LYSINE BIOSYNTHESIS; RESISTANT MUTANTS; SPLIT PATHWAY; GENES; AMPLIFICATION; FERMENTATION**

L15 ANSWER 9 OF 13 CAPLUS COPYRIGHT 2001 ACS  
ACCESSION NUMBER: 1985:436156 CAPLUS  
DOCUMENT NUMBER: 103:36156  
TITLE: Optimization of amino acid production by automatic self-tuning digital control of redox potential  
AUTHOR(S): Radjai, Mohammad K.; Hatch, Randolph T.; Cadman, Theodore W.  
CORPORATE SOURCE: Dep. Chem. Nucl. Eng., Univ. Maryland, College Park, MD, 20742, USA  
SOURCE: Biotechnol. Bioeng. Symp. (1984), 14(Symp. Fuels Chem., 6th), 657-79  
DOCUMENT TYPE: CODEN: BIBSBR; ISSN: 0572-6565  
LANGUAGE: Journal English  
AB The **microbial prodn.** of homoserine [672-15-1], **lysine** [56-87-1], and valine [72-18-4] by an auxotrophic mutant of **Corynebacterium glutamicum** was investigated in a 16-L batch fermentor. Closed-loop digital control of redox potential was implemented using proportional-integral (PI) control of agitation rates. Due to the nonlinearity of the system, the PI controller parameters had to be changed during the course of the fermns. An automatic, self-tuning algorithm was developed for stable control of redox potential. This permitted exptl. optimization of total and selective amino acid prodn. Total amino acid yields of 35% from glucose were achieved compared to 23% reported in the literature for the same fermn.  
ST amino acid fermn redox potential control; optimization simulation

homoserine lysine valine fermn  
IT **Corynebacterium glutamicum**  
(amino acid manuf. with, optimization and redox potential control in)  
IT **Amino acids, preparation**  
RL: BMF (Bioindustrial manufacture); BIOL (Biological study); PREP  
(Preparation)  
(manuf. of, by fermn.)

L15 ANSWER 10 OF 13 MEDLINE  
ACCESSION NUMBER: 79079819 MEDLINE  
DOCUMENT NUMBER: 79079819 PubMed ID: 727028  
TITLE: **Microbial production** of essential amino acid with **Corynebacterium glutamicum** mutants.  
AUTHOR: Nakayama K; Araki K; Kase H  
SOURCE: ADVANCES IN EXPERIMENTAL MEDICINE AND BIOLOGY, (1978) 105 649-61.  
PUB. COUNTRY: Journal code: 2LU; 0121103. ISSN: 0065-2598.  
United States  
Journal; Article; (JOURNAL ARTICLE)  
LANGUAGE: English  
FILE SEGMENT: Priority Journals  
ENTRY MONTH: 197902  
ENTRY DATE: Entered STN: 19900314  
Last Updated on STN: 19970203  
Entered Medline: 19790212

TI **Microbial production** of essential amino acid with **Corynebacterium glutamicum** mutants.  
AB **Amino acids** produced by microbial process are generally L-forms. The stereospecificity of the **amino acids** produced by fermentation makes the process advantageous compared with synthetic process. Microorganisms employed in microbial process for amino acid production are divided into 4 classes; wild-type strain, auxotrophic mutant, regulatory mutant and auxotrophic regulatory mutant. Using such mutants of **Corynebacterium glutamicum**, all the essential **amino acids** but L-methionine are now being produced by "direct fermentation" from cheap carbon sources such as carbohydrate materials or acetic acid.

CT \***Amino Acids, Essential: BI, biosynthesis**  
\***Corynebacterium: ME, metabolism**  
Fermentation  
Kinetics  
Leucine: BI, biosynthesis  
**Lysine: BI, biosynthesis**  
Mutation  
Phenylalanine: BI, biosynthesis  
Species Specificity  
Stereoisomerism  
Threonine: BI, biosynthesis  
Tryptophan: BI, biosynthesis

RN 3617-44-5 (Phenylalanine); 56-87-1 (**Lysine**); 7005-03-0  
(Leucine); 72-19-5 (Threonine); 73-22-3 (Tryptophan)

CN 0 (**Amino Acids, Essential**)

L15 ANSWER 11 OF 13 CAPLUS COPYRIGHT 2001 ACS  
ACCESSION NUMBER: 1970:508239 CAPLUS  
DOCUMENT NUMBER: 73:108239  
TITLE: **Microbial production** of L-threonine

INVENTOR(S): Nakayama, Kiyoshi; Kase, Hiroshi  
PATENT ASSIGNEE(S): Kyowa Fermentation Industry Co. Ltd.  
SOURCE: Ger. Offen., 22 pp.

CODEN: GWXXBX

DOCUMENT TYPE: Patent

LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
DE 1817666	A	19700827	DE 1968-1817666	19681224

TI **Microbial production of L-threonine**

AB Various microorganisms, e.g. Aerobacter [Enterobacter] aerogenes, Serratia

marcescens, or Arthrobacter paraffineus, cultured for producing L-threonine required 2 or 3 of the **amino acids** isoleucine, methionine, **lysine**, or diaminopimelic acid. The microorganisms were cultured aerobically in an aq. medium contg. the optimal (or less) amts. of the required **amino acids**. Thus, E. aerogenes NM-IS-5 (ATCC 21,215) was cultured 96 hr at 30.degree. in medium contg. glucose 5, (NH4)2SO4 1.4, KH2PO4 0.05, K2HPO4 0.05, MgSO4·7H2O 0.025, FeSO4·7H2O 0.001, MnSO4·4H2O 0.001, and CaCO3 2% and isoleucine 50, methionine 100, and diaminopimelic acid 200 mg/l. to give 7.8 g L-threonine/l.

ST **microbial prodn threonine; threonine microbial**

**prodn; Aerobacter threonine fermn; amino acid prodn fermn**

IT **Corynebacterium**

(glutamicum, threonine manuf. by)

L15 ANSWER 12 OF 13 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1967:514494 CAPLUS

DOCUMENT NUMBER: 67:114494

TITLE:

**Microbial production of amino acids from hydrocarbons. III.**

L-Ornithine production by an arginine auxotrophic mutant of **Corynebacterium** hydrocarboclastus

AUTHOR(S): Ishu, Ryosuke; Ishii, Ryosuke; Shio, Isamu

CORPORATE SOURCE: Ajinomoto Co., Inc., Kawasaki, Japan

SOURCE: J. Gen. Appl. Microbiol. (1967), 13(3), 3303-12

CODEN: JGAMA9

DOCUMENT TYPE: Journal

LANGUAGE: English

TI **Microbial production of amino acids**

from hydrocarbons. III. L-Ornithine production by an arginine auxotrophic mutant of **Corynebacterium** hydrocarboclastus

AB cf. CA 67: 89718u. The arginine auxotrophic mutant strain RN-362 of C. hydrocarboclastus R-7 was used to study L-ornithine production from hydrocarbons, in a fermentation medium contg. various n-alkanes. L-Ornithine production required L-arginine at the optimum level of 0.5-1.0

g./l. of medium; an excess inhibited the biosynthesis of L-ornithine. (NH4)2HPO4 was the best source of N and, at 2% in a neutral to slightly acidic pH, gave the highest level of L-ornithine production and cell growth; NH4OAc, KNO3, and (NH4)2CO3 proved less suitable because of a drop

in pH along with the accumulation of a large amt. of .alpha.-ketoglutaric acid, pyruvic acid, and proline in the growth medium. Of 17 C sources,

n-tetradecane best supported cell growth and L-ornithine production and the other C13-C17 n-alkanes did so moderately, while kerosene and light oil produced good cell growth but only a small amt. of L-ornithine.

Addn.

of 3 g. yeast ext. and 0.5 g. L-arginine-HCl to 1 l. of medium enhanced L-ornithine production. A similar effect was achieved by replacing the yeast ext. with various **amino acids** at 0.01% in the medium. L-Methionine was most effective for the production of L-ornithine, while L-lysine, L-cysteine, L-cystine, L-histidine, and L-phenylalanine were less so, in decreasing order. **Amino acids** enhance L-ornithine production by stimulating hydrocarbon oxidn. and cell growth.

ST HYDROCARBONS USE BACTERIA; BACTERIA HYDROCARBONS USE; ALKANES USE BACTERIA; **AMINO ACIDS** PRODN HYDROCARBONS; ORNITHINE PRODN HYDROCARBONS; PARAFFINS UTILIZATION BACTERIA

IT **Corynebacterium**

(hydrocarboclastus, ornithine formation from hydrocarbons by)

IT Hydrocarbons, biological studies

RL: BIOL (Biological study)

(ornithine formation from, by **Corynebacterium** hydrocarboclastus)

IT 70-26-8

RL: FORM (Formation, nonpreparative)

(formation of, from hydrocarbons by **Corynebacterium** hydrocarboclastus)

L15 ANSWER 13 OF 13 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1967:489718 CAPLUS

DOCUMENT NUMBER: 67:89718

TITLE:

**Microbial production of amino acids from hydrocarbons. II.**

Isolationf good hycarbon utilizers and amino acid production by their auxotrophs

AUTHOR(S): Ishii, Ryosuke; Otsuka, Shinichiro; Shio, Isamu

CORPORATE SOURCE: Central Res. Labs., Ajinomoto Co., Inc., Kawasaki, Japan

SOURCE:

J. Gen. Appl. Microbiol. (1967), 13(2), 217-25

CODEN: JGAMA9

DOCUMENT TYPE: Journal

LANGUAGE: English

TI **Microbial production of amino acids**

from hydrocarbons. II. Isolationf good hycarbon utilizers and amino acid

production by their auxotrophs

AB cf. CA 59: 14313h. Nine microorganisms, which showed good growth on long-chain aliphatic hydrocarbons, were isolated by an enrichment culture method, followed by a single colony isolation technique. They included 5 strains of Alcaligenes marshallii, 2 strains of **Corynebacterium** hydrocarboclastus, and 2 strains of yeast. Various auxotrophic mutants were derived from these microorganisms. The mutants accumulated the following **amino acids** from aliphatic hydrocarbons; L-ornithine, L-valine, L-glutamic acid, L-leucine, L-tyrosine, L-alanine, L-proline, L-aspartic acid, and L-lysine.

ST BACTERIA AMINO ACID PRODN; AMINO ACID PRODN BACTERIA; HYDROCARBONS

**AMINO ACIDS; ALIPHATICS BACTERIA METAB**

IT **Corynebacterium**

(hydrocarboclastus, amino acid fermentation of hydrocarbons by)

IT **Amino acids, preparation**

RL: BMF (Bioindustrial manufacture); BIOL (Biological study); PREP  
(Preparation)  
(manuf. of, by fermentation of hydrocarbons)

=> dis his

(FILE 'HOME' ENTERED AT 18:56:31 ON 19 OCT 2001)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, SCISEARCH' ENTERED AT 18:57:06 ON  
19 OCT 2001

L1        2893 S MICROBIAL (W) PRODUCTION  
L2        190 S L1 AND (AMINO (W) ACIDS)  
L3        28 S L2 AND CORYNEBACTERIUM  
L4        13 L3 AND LYSINE  
L5        0 L3 AND ((EXPORT) (W) (GENE OR CARRIER))  
L6        0 S L3 AND EXPORT (W) GENE  
L7        166 S EXPORT (W) GENE  
L8        0 S L3 AND L7  
L9        0 S L3 (P) L7  
L10      0 S L2 AND L7  
L11      0 S L2 AND EXPORT (W) GENE  
L12      61 S L7 AND MICROB?  
L13      1 S L12 AND CORYNEBACTERIUM  
L14      26 DUP REM L3 (2 DUPLICATES REMOVED)  
L15      13 DUP REM L4 (0 DUPLICATES REMOVED)

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\* \* \* \* \*

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STN INTERNATIONAL LOGOFF AT 19:32:29 ON 19 OCT 2001